Appl. No. 09/438,358

## (12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

# (19) World Intellectual Property Organization International Bureau



## 

### (43) International Publication Date 1 February 2001 (01.02.2001)

### PCT

# (10) International Publication Number WO 01/07572 A3

(51) International Patent Classification?: 15/79, 15/90 // 9/22

C12N 5/10,

- (21) International Application Number: PCT/US00/19983
- (22) International Filing Date: 21 July 2000 (21.07.2000)
- (25) Filing Language:

**English** 

(26) Publication Language:

English

(30) Priority Data: 60/145,469

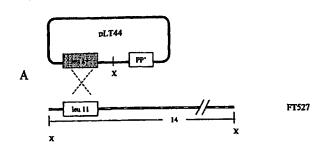
23 July 1999 (23.07.1999) US

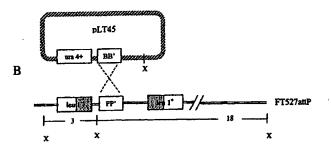
(71) Applicants: THE REGENTS OF THE UNIVERSITY OF CALIFORNIA [US/US]; Office Of Technology Transfer, 1111 Franklin Street, 12th floor, Oakland, CA 94607-5200 (US). THE UNITED STATES OF AMERICA as represented by the SECRETARY OF AGRICULTURE [US/US]; 1400 Independence Avenue S.W., Washington, DC 20250 (US).

- (72) Inventors: OW, David, W.; 127 Brenner Street, Hercules, CA 94547-3749 (US). CALENDAR, Richard; 940 Euclid Street, Berkeley, CA 94708-1436 (US). THOMASON, Lynn; 1061 H. Monroe Street #7, Albany, CA 94706-2282 (US).
- (74) Agents: SMITH, Timothy, L. et al.; Townsend and Townsend and Crew LLP, Two Embarcadero Center, 8th floor, San Francisco, CA 94111-3834 (US).
- (81) Designated States (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW.
- (84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW), Eurasian

[Continued on next page]

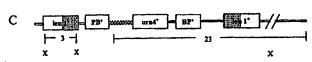
(54) Title: DNA RECOMBINATION IN EUKARYOTIC CELLS BY THE BACTERIOPHAGE PHIC31 RECOMBINATION SYSTEM





(57) Abstract: This invention provides methods for obtaining specific and stable integration of nucleic acids into eukaryotic cells. The invention makes use of site-specific recombination systems that use prokaryotic recombinase polypeptides, such as the  $\Phi$ C31 integrase, that can mediate recombination between the recombination sites, but not between hybrid recombination sites that are formed upon the recombination. Thus, the recombination is irreversible in the absence of additional factors. Eukaryotic cells that contain the recombinase polypeptides, or genes that encode the recombinases, are also provided.





## WO 01/07572 A3



patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

#### Published:

With international search report.

(88) Date of publication of the international search report: 19 April 2001

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12N5/10 C12N //C12N9/22 C12N15/90 C12N15/79 According to International Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) C12N IPC 7 Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) EPO-Internal, WPI Data, PAJ, BIOSIS, CHEM ABS Data, EMBASE C. DOCUMENTS CONSIDERED TO BE RELEVANT Relevant to dalm No. Citation of document, with indication, where appropriate, of the relevant passages 1-8, WO 00 11155 A (UNIV LELAND STANFORD P.X 10-47 JUNIOR) 2 March 2000 (2000-03-02) page 4 page 5, line 25 -page 6, line 2 1-47 THORPE, H. M. ET AL.: "In vitro A site-specific integration of bacteriophage DNA catalyzed by a recombinase of the resolvase/invertase family" PNAS, vol. 95, 1998, pages 5505-5510, XP002157128 cited in the application abstract Patent family members are listed in annex. Further documents are listed in the continuation of box C. X "I" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "E" earlier document but published on or after the international filing date \*L\* document which may throw doubts on priority claim(s) or which is clied to establish the publication date of another citation or other special reason (as specified) "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled by the cart. "O" document referring to an oral disclosure, use, exhibition or other means document published prior to the international filing date but later than the priority date claimed \*&\* document member of the same patent family Date of mailing of the international search report Date of the actual completion of the international search 29/01/2001 12 January 2001 Authorized officer Name and mailing address of the ISA

Fax: (+31-70) 340-3016

1

European Patent Office, P.B. 5818 Patentiaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo ni,

Mata Vicente, T.

#### TIONAL SEARCH REPORT INTER information on patent family members

nd Application No PCT/US 00/19983

Patent document cited in search report		Publication date	Patent family member(s)		Publication date
WO 0011155	A	02-03-2000	AU	5898599 A	14-03-2000
WO 9918222	A	15-04-1999	AU EP	9466898 A 1021550 A	27-04-1999 26-07-2000